Conclusion

No fees are believed due; however, should any fees be required, please charge them to Deposit Account No. 11-1410. A duplicate copy of this communication is enclosed. Should there be any questions concerning this application, the Examiner is respectfully invited to contact the undersigned at the telephone number appearing below.

Respectfully submitted,

KNOBBE, MARTENS, OLSON & BEAR, LLP

Dated: 10 May 2001

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VERSION WITH MARKINGS TO SHOW CHANGES MADE

IN THE SPECIFICATION

On page 44, 5th paragraph (lines 24-30), please replace the paragraph with the following paragraph: --Plasmid pCR.Bgl-GFP-Bam (Figure 5) comprises an internal region of the GFP open reading frame derived from plasmid pEGFP-N1 MCS (Figure 1) placed operably under the control of the lacZ promoter. To produce this plasmid, a region of the GFP open reading frame was amplified from pEGFP-N1 MCS using the amplification primers Bgl-GFP (SEQ ID NO:1) and GFP-*Bam* (SEQ ID NO:2) and cloned into plasmid pCR2.1. The internal GFP-encoding region in plasmid pCR.Bgl-GFP-Bam lacks functional translational start and stop codons.--.

On page 45, 3rd paragraph (lines 15-21), please replace the paragraph with the following paragraph: --Plasmid pCR.SV40L (figure 8) comprises the SV40 late promoter derived from plasmid pSVL (GenBank Accession No. U13868; Pharmacia), cloned into pCR2.1 (Stratagene). To produce this plasmid, the SV40 late promoter was amplified using the primers SV40-1 (SEQ ID NO:3) and SV40-2 (SEQ ID NO:4) which comprise *Sal* I cloning sites to facilitate sub-cloning of the amplified DNA fragment into pCMV.cass. The primer also contains a synthetic poly (A) site at the 5' end, such that the **[amplicification]**amplification product comprises the synthetic poly(A) site at the 5' end of the SV40 promoter sequence.--.

On page 45 4th paragraph (line 24) through page 46, (line 5), please replace the paragraph with the following: --The BEV RNA-dependent RNA polymerase coding region was amplified as a 1,385 bp DNA fragment from a full-length cDNA clone encoding same, using primers designated BEV-1 (SEQ ID NO:5) and BEV-2 (SEQ ID NO:6), under standard amplification conditions. The amplified DNA contained a 5'-*Bgl* II restriction enzyme site, derived from the BEV-1 primer sequence (SEQ ID NO:5) and a 3'*Bam*HI restriction enzyme site, derived from the BEV-2 primer sequence (SEQ ID NO:6). Additionally, as the BEV-1 primer sequence (SEQ ID NO:5) contains a translation start signal 5'-ATG-3' engineered at positions 15-17, the amplified BEV polymerase structural gene comprises the start site in-frame with BEV polymerase-encoding nucleotide sequences, Thus, the amplified BEV polymerase structural gene comprises the ATG start codon immediately upstream (ie. Juxtaposed) to the BEV polymerase-encoding sequence. There is no translation stop codon in the amplified DNA. This plasmid is present as Figure 9.--.

On page 46, 2nd paragraph (lines 17-25), please replace the paragraph with the following:

--A non-translatable BEV polymerase structural gene was amplified from a full-length BEV polymerase cDNA clone using the amplification primers BEV-3 (SEQ ID NO:7) and BEV-4(SEQ ID NO:8). Primer BEV-4 comprises a *Bgl*II cloning site at positions 5-10 and sequences downstream of this *Bgl*II site are homologous to nucleotide sequences of the BEV polymerase gene. There is no functional ATG start codon in the amplified DNA product of primers BEV-3 (SEQ ID NO:7) and BEV-4 (SEQ ID NO:8). The BEV polymerase is expressed as part of a polyprotein and, as a consequence, there is no ATG translation start site in this gene. The amplified DNA was cloned into plasmid pR2.1 (Stratagene) to yield plasmid pCR.BEV.3 (Figure 11).

On page 57, paragraphs 5 and 6 (lines 12-16), please replace the paragraph with the following:

--NOS 5' (forward primer; SEQ ID [??] NO:9)

5'-GGATTCCCGGGACGTCGCGAATTTCCCCCGATCGTTC-3'; and

NOS 3' (reverse primer; SEQ ID [??] NO:10)

5'-CCATGGCCATATAGGCCCGATCTAGTAACATAG-3'--.

On page 60, paragraph 3 (lines 9-11), please replace the paragraph with the following:

--LNYV 1:5' ATGGGATCCGTTATGCCAAGAAGAAGGA-3'(SEQ ID NO:11); and

LNYV 2:5' ATGGGATCCGTTATGCCAAGAAGAAGGA-3'(SEQ ID NO:12)--.

On page 66, paragraph 2 (lines 9-15), please replace the paragraph with the following:

--PVY1:

5'TAATGAGGATGATGTCCCTACCTTTAATTGGCAGAAATTTCTGTGGAAAGACAGGG AAATCTTTCGGCATTT-3'(SEQ ID NO:13); and

PVY 2:

5' TTCTGCCAATTAAAGGTAGGGACATCATCCTCATTAAAATGCCGAAAGATTTCCCT GTCTTTCCACAGAAAT-3'(SEQ ID NO:14)--.